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C:\Program Files\Stnexp\Queries\10694561
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15-16 16-17 17-18
exact/norm bonds:
    1-2 1-6 1-20 2-3 2-11 3-4 4-5 4-10 5-6 5-7 6-9 7-8 8-9 11-12
    21-22 21-23 23-24
exact bonds:
    12-17 20-21
normalized bonds:
    13-14 13-18 14-15 15-16 16-17 17-18
isolated ring systems:
    containing 1:

Match level:
    1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom
    10:CLASS 11:CLASS 12:CLASS 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom
    18:Atom 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS
```

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 13-14 13-18 14-15

chain nodes :

ring nodes :

ring bonds :

chain bonds :

10 11 12

1 2 3 4

20 21

6

5

22

1-20 2-11 4-10 11-12 12-17 20-21

23 24

7 8 9 13 14 15 16 17

18

21-22 21-23 23-24

=>

Uploading C:\Program Files\Stnexp\Queries\10694561.str

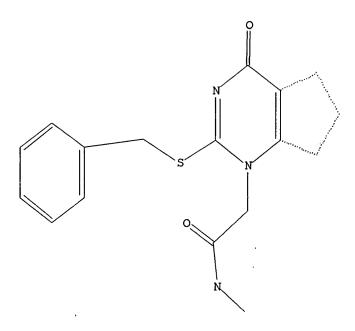
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chain nodes :
10 11 12 20 21 22 23 24
ring nodes :
             6 7 8 9 13 14 15 16 17 18
chain bonds :
1-20 2-11 4-10 11-12 12-17 20-21 21-22 21-23 23-24
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 13-14 13-18 14-15 15-16 16-17
17-18
exact/norm bonds :
1-2 1-6 1-20 2-3 2-11 3-4 4-5 4-10 5-6 5-7 6-9 7-8 8-9 11-12 21-22
21-23 23-24
exact bonds :
12-17 20-21
normalized bonds :
13-14 13-18 14-15 15-16 16-17 17-18
isolated ring systems :
containing 1:
```

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS 11:CLASS 12:CLASS 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS

L1 STRUCTURE UPLOADED

=> d l1 L1 HAS NO ANSWERS L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11 sss sam
SAMPLE SEARCH INITIATED 20:11:09 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 34 TO ITERATE

100.0% PROCESSED 34 ITERATIONS 1 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 331 TO 1029
PROJECTED ANSWERS: 1 TO 80

L2 1 SEA SSS SAM L1

=> s l1 sss ful FULL SEARCH INITIATED 20:11:17 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 612 TO ITERATE

100.0% PROCESSED 612 ITERATIONS 29 ANSWERS SEARCH TIME: 00.00.01

L3 29 SEA SSS FUL L1

=> => s 13 L4 6 L3

=> d 14 1-6 bib, ab, hitstr

- L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2004:720873 CAPLUS
- DN 141:342698
- TI SB-480848 (GlaxoSmithKline)
- AU Rotella, David P.
- CS Lexicon Pharmaceuticals, Princeton, NJ, 08540, USA
- SO Current Opinion in Investigational Drugs (Thomson Scientific) (2004) 5(3), 348-351 CODEN: COIDAZ; ISSN: 1472-4472
- PB Thomson Scientific
- DT Journal; General Review
- LA English
- AB A review. SB-480848 (synonyms/analogs: SB-435445, Lp-PLA2 inhibitor) is a reversible lipoprotein-associated phospholipase A2 inhibitor under development by GlaxoSmithKline for the potential treatment of atherosclerosis. Phase II trials with SB-480848 are currently underway.
- IT 356057-34-6P, SB 480848
 RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(SB 435445; reversible lipoprotein-associated phospholipase A2 inhibitor SB-480848 for potential treatment of atherosclerosis)

- RN 356057-34-6 CAPLUS
- CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
ANSWER 2 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
L4
     2003:837075 CAPLUS
AN
DN
     139:337982
ΤI
     Preparation of pyridone and pyrimidone compounds as inhibitors of the
     enzyme Lp-PLA2
IN
     Leach, Colin Andrew; Smith, Stephen Allan
PA
     Glaxo Group Limited, UK
SO
     PCT Int. Appl., 61 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
                                   ÁTE
     PATENT NO.
                           KIND
                                               APPLICATION NO.
                                                                        DATE
                                  20031023
                                               WO 2003-GB1550
PΙ
     WO 2003087088
                            A2
                                                                        20030410
     WO 2003087088
                                  20040108
                            Α3
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              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
              PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
              UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
              BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI GB 2002-8280
                                  20020410
                            Α
os
     MARPAT 139:337982
AB
     The title compds. [I; R1 = (un)substituted aryl; R2 = halo, alkyl, alkoxy,
     etc.; R3 = H, halo, alkyl, hydroxyalkyl; R2 and R3 together with the
     pyridone or pyrimidine ring carbons to which they are attached form
     (un) substituted fused 5-6 membered carbocyclic ring, fused benzo or
     heteroaryl ring; R4 = (CH2)n substituted by benzimidazole or 5-6 membered
     heteroaryl; R5 = (un)substituted (hetero)aryl; R6 = (un)substituted
     (hetero)aryl; X = CH, N; Y = alkylene, CH: CH: CH2)mS; n = 1-4; m = 1-2
     that are inhibitors of the enzyme Lp-PLA2 and are of use in therapy, in
     particular for treating atherosclerosis, were prepared Thus, amidation of
     2-[2-(2,3-difluorobenzylthio)-4-oxo-4H-quinolin-1-yl]acetic acid with
     N-[2-(1-methylimidazol-4-yl)ethyl]-4'-trifluoromethylbiphen-4-
     ylmethylamine (prepns. given) afforded the quinolinone II. The
     exemplified compds. I showed IC50 values in the range <0.1 to 100 nM
     against Lp-PLA2.
IT
     615578-19-3P
     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
         (preparation of pyridone compds. as inhibitors of the enzyme Lp-PLA2)
RN
     615578-19-3 CAPLUS
     1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-
CN
     4,5,6,7-\text{tetrahydro-N-}[2-(1-\text{methyl-1H-imidazol-4-yl})\text{ ethyl}]-4-\text{oxo-N-}[[4'-
```

(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

CH₂-s

PAGE 2-A

```
ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
L4
ΑN
     2003:836853 CAPLUS
DN
     139:337978
ΤI
     Preparation of N-substituted pyridinone and pyrimidinone derivatives for
     use as Lp-PLA2 inhibitors in the treatment of atherosclerosis
IN
     Leach, Colin Andrew; Smith, Stephen Allan
PA
     Glaxo Group Limited, UK
SO
     PCT Int. Appl., 38 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                           KIND
                                               APPLICATION NO.
                                                                        DATE
                                  20031023
                                               WO 2003-GB1544
                                                                        20030410
PΙ
     WO 2003086400
                            A1
         W: AE, AG, AL, AM, AT\ AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
              PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
         UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
              FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI GB 2002-8279
                                  20020410
     MARPAT 139:337978
os
AΒ
     The title compds. [I; R1 = (un)substituted aryl; R2 = halo, alkyl, alkoxy,
     etc.; R3 = H, halo, alkyl, hydroxyalkyl; R2 and R3 together with the
     pyridone or pyrimidone ring carbons to which they are attached form
     (un) substituted fused 5-6 membered carbocyclic ring, fused benzo or
     heteroaryl ring; R4 = alkyl substituted by 5-7 membered saturated heterocyclyl
     comprising N and optionally O or S; R5 = (un)substituted (hetero)aryl; R6
     = (un)substituted (hetero)aryl; X = CH, N; Y = alkylene, CH:CH, (CH2)nS; n
     = 1-3] that are inhibitors of the enzyme Lp-PLA2 and are of use in
     therapy, in particular for treating atherosclerosis, were prepared Thus,
     amidation of 2-[2-(2,3-difluorobenzylthio)-4-oxo-4H-quinolin-1-yl]acetic
     acid with N-(1-thiazol-2-ylmethylpiperidin-4-yl)-4'-trifluoromethylbiphen-
     4-ylmethylamine (prepns. given) afforded the quinolinone II. The
     exemplified compds. I showed IC50 values in the range <0.1 to 100 nM
     against Lp-PLA2.
IT
     615577-22-5P
     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
         (preparation of pyridinone and pyrimidinone derivs. for use as Lp-PLA2
        inhibitors in the treatment of atherosclerosis)
     615577-22-5 CAPLUS
RN
     1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-
     4,5,6,7-tetrahydro-4-oxo-N-[1-(phenylmethyl)-4-piperidinyl]-N-[[4'-
```

(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 2-A

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/694,561

- L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2003:215748 CAPLUS
- DN 139:78433
- TI The identification of clinical candidate SB-480848: a potent inhibitor of lipoprotein-associated phospholipase A2
- AU Blackie, Josie A.; Bloomer, Jackie C.; Brown, Murray J. B.; Cheng, Hung-Yuan; Hammond, Beverley; Hickey, Deirdre M. B.; Ife, Robert J.; Leach, Colin A.; Lewis, V. Ann; Macphee, Colin H.; Milliner, Kevin J.; Moores, Kitty E.; Pinto, Ivan L.; Smith, Stephen A.; Stansfield, Ian G.; Stanway, Steven J.; Taylor, Maxine A.; Theobald, Colin J.
- CS Medicines Research Centre, GlaxoSmithKlide, Stevenage, SG1 2NY, UK
- SO Bioorganic & Medicinal Chemistry Letters (2003), 13(6), 1067-1070 CODEN: BMCLE8; ISSN: 0960-894X
- PB Elsevier Science B.V.
- DT Journal
- LA English
- OS CASREACT 139:78433
- AB Modification of the pyrimidone 5-substituent in clin. candidate SB-435495 has given a series of inhibitors of recombinant lipoprotein-associated phospholipase A2 with sub-nanomolar potency. Cyclopentyl fused derivative 21, SB-480848, showed an enhanced in vitro and in vivo profile vs. SB-435495 and has been selected for progression to man.
- IT 552857-62-2P 552857-63-3P
 - RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 - (design and structure activity of lipoprotein-associated phospholipase A2 inhibitor SB-480848)
- RN 552857-62-2 CAPLUS
- CN 1H-Cyclopentapyrimidine-1-acetamide, N-(2-aminoethyl)-N-[(4'-chloro[1,1'-biphenyl]-4-yl)methyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-(9CI) (CA INDEX NAME)

PAGE 2-A

RN 552857-63-3 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-(2-aminoethyl)-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

CF3
$$\begin{array}{c} CH_2 \\ N-CH_2-CH_2-NH_2 \\ C=0 \\ CH_2 \end{array}$$

PAGE 2-A

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L4
     ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     2003:154411 CAPLUS
     138:187787
DN
     Novel processes for the preparation of pyrimidinone derivatives, useful as
ΤI
     Lp-PLA2 inhibitors, and intermediates thereof
IN
     Mulholland, Keith Raymond; Ross, Andrew R.; Slater, Graham Ralph; Smith,
     Gillian Elizabeth
PA
     Smithkline Beecham PLC, UK
                                                              Common Assignee
SO
     PCT Int. Appl., 14 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                                            APPLICATION NO.
                         KIND
                                DATE
                                                                    DATE
                                _____ ·
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                         ____
                                            -----
    WO 2003016287
                         A2
                                20030227
                                            WO 2002-EP9067
                                                                    20020813
PΙ
    WO 2003016287 '
                         A3
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             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF,
             CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                20040915 EP 2002-794787
     EP 1456183
                          A2
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
                                                                               Queck
                                20041202
                                           US 2004-485972
                                                                    20040702
     US 2004242875
                         A1
PRAI GB 2001-19795
                          Α
                                20010814
     WO 2002-EP9067
                          W
                                20020813
OS
     CASREACT 138:187787; MARPAT 138:187787
AΒ
     The invention relates to a process for the preparation of certain pyrimidinone
     compds., including the intermediates I and II [wherein: RaRb = (CH2)3-4;
     R1 = Ph optionally substituted by halogen], and the final target compds.
     III [wherein: RaRb = atoms to form 5-membered carbocyclic ring; R1 =
     4-fluorophenyl; R2 = C1-3 alkyl substituted by NR5R6; R2 = Het-C0-2-alkyl;
     Het = 5- to 7-membered N-heterocyclyl with N optionally substituted by
     C1-6 alkyl; R3R4 = 4-[4-(trifluoromethyl)phenyl]phenyl; R5, R6 = H, C1-6
     alkyl]. Compds. III, described in WO 01/60805, are known inhibitors (no
     data) of lipoprotein-associated phospholipase A2 (Lp-PLA2), useful, e.g., for
    prevention of acute coronary events caused by atherosclerosis. The
     literature methods of preparing III suffer from moderate yields due to poor
     selectivity in the alkylation of the pyrimidinone nucleus. The invention
    method gives selective N1-alkylation of the pyrimidinone nucleus, and does
    not require isolation of an intermediate ester, thus giving high yields
     and efficiency. For instance, cyclocondensation of Et
     2-oxocyclopentanecarboxylate with thiourea in the presence of DBU gave
     67.6% 5,6-trimethylene-2-thiouracil, which underwent S-alkylation by
     4-fluorobenzyl chloride in the presence of K2CO3 and KI in Me2CO to give
     86.5% intermediate II [R1 = 4-FC6H4, RaRb = (CH2)3]. This compound was
     selectively O-silylated by (Me3Si)2NH and saccharin in CH2Cl2, selectively
     N1-alkylated by CF3SO2OCH2CO2Me, and then hydrolyzed directly by aqueous NaOH
     in iso-PrOH, to give 69% I [R1 = 4-FC6H4, RaRb = (CH2)3]. Amidation of
     this acid with the corresponding amine using DIPEA and TBTU in CH2Cl2,
     followed by recrystn. from iso-PrOAc, gave 88% target compound IV.
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PAGE 1-A

CH2-S-II

PAGE 2-A

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L4
     ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
AN
     2001:617985 CAPLUS
     135:195570
DN
     Preparation of pyrimidine-4-one derivatives as LDL-PLA2 inhibitors
ΤI
     Hickey, Deirdre Mary Bernadette; Ife, Robert John; Leach, Colin Andrew;
IN
     Pinto, Ivan Leo; Smith, Stephen Allan; Stanway, Steven James
     Smithkline Beecham P.L.C., UK
PA
SO
     PCT Int. Appl., 54 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
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PΙ
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             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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                           AA
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                                 20010827
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                                 20021211
                                              EP 2001-907522
                                                                      20010213
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                           Α
                                              JP 2001-560190
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                           T2
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     NZ 520752
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                                 20040326
                                              NZ 2001-520752
                                                                      20010213
     RU 2235722
                           C2
                                 20040910
                                              RU 2002-124611
                                                                      20010213
     US 2002103213
                           A1
                                 20020801
                                              US 2001-782930
                                                                      20010214
                           В
                                 20030901
                                              TW 2001-90103332
                                                                      20010215
     TW 550259
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     NO 2002003828
                           Α
                                              NO 2002-3828
                                                                      20020813
     ZA 2002006528
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                                 20030313
                                              ZA 2002-6528
                                                                      20020815
     BG 107034
                           Α
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                                              BG 2002-107034
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     US 6649619
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                                                                      20030203
                           В1
     US 2004097525
                           Α1
                                 20040520
                                              US 2003-694561
                                                                      20031027
PRAI GB 2000-3636
                           Α
                                 20000216
     GB 2001-1437
                           Α
                                 20010119
     WO 2001-EP1515
                           W
                                 20010213
     US 2001-782930
                                 20010214
                           В1
                                 20030203
     US 2003-357238
                           A3
OS
     MARPAT 135:195570
     The title compds. [I; Ra = H, halo, alkyl, etc.; Rb = H, halo, alkyl,
AB
     etc.; Ra and Rb together = (CH2)n (n = 3-4) or Ra and Rb together with the
     pyrimidine ring carbon atoms to which they are attached form
     (un) substituted fused benzo or heteroaryl ring; Rc = H, alkyl; R2 =
     (un) substituted (hetero) aryl; R3 = H, alkyl, halo, etc.; R4 =
     (un) substituted (hetero) arylene; R5 = (un) substituted (hetero) aryl; n =
     1-4; X = 0, S; Y = (CH2)pOq (p = 1-3 and q = 0; p = 2-3 and q = 1); Z = 0,
     a bond] which are inhibitors of the enzyme Lp-PLA2 useful in treating atherosclerosis, were prepared Thus, reacting N-[2-(diethylamino)ethyl]-4-
     (4-trifluoromethylphenyl)benzylamine with 1-(carboxymethyl)-2-(4-
     fluorobenzylthio)-5-ethylpyrimidin-4-one in the presence of HATU and
     (iso-Pr)2NEt in CH2Cl2 afforded the pyrimidinone II. The compds. I
```

described in Examples were tested for Lp-PLA2 inhibition and showed IC50 values in the range <0.1 nM to 10 μM .

IT 356057-38-0P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of pyrimidine-4-one derivs. as LDL-PLA2 inhibitors)

RN 356057-38-0 CAPLUS

1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[6-[4-(trifluoromethyl)phenyl]-3-pyridinyl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

CN

CRN 356057-37-9 CMF C35 H37 F4 N5 O2 S

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CM 2
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CRN 87-69-4 CMF C4 H6 O6

Absolute stereochemistry.

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356057-34-6P 356057-35-7P 356057-36-8P
IT
     356057-37-9P 356057-39-1P 356057-40-4P
     356057-69-7P 356057-87-9P 356057-88-0P
     356057-89-1P 356057-90-4P 356057-91-5P
     356057-92-6P 356057-93-7P 356057-94-8P
     356057-95-9P 356057-98-2P 356057-99-3P
     356058-00-9P 356058-03-2P 356058-05-4P
     356058-06-5P 356058-07-6P 356058-12-3P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of pyrimidine-4-one derivs. as LDL-PLA2 inhibitors)
RN
     356057-34-6 CAPLUS
CN
     1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-
     fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-
     (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)
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RN 356057-35-7 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 356057-34-6 CMF C36 H38 F4 N4 O2 S

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CM 2
CRN 87-69-4
CMF C4 H6 O6

Absolute stereochemistry.

RN 356057-36-8 CAPLUS CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-

fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-, monohydrochloride (9CI)
(CA INDEX NAME)

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HC1

RN 356057-37-9 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[6-[4-(trifluoromethyl)phenyl]-3-pyridinyl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-39-1 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[2-[4-(trifluoromethyl)phenyl]-5-pyrimidinyl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-40-4 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[2-[4-(trifluoromethyl)phenyl]-5-pyrimidinyl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 356057-39-1 CMF C34 H36 F4 N6 O2 S

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CM 2
CRN 87-69-4
CMF C4 H6 O6

Absolute stereochemistry.

RN 356057-69-7 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[(4'-chloro[1,1'-biphenyl]-4-

yl)methyl]-N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio}-4,5,6,7-tetrahydro-4-oxo-(9CI) (CA INDEX NAME)

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RN 356057-87-9 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(2,3-difluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-88-0 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(3,4-difluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-89-1 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-2-[[(2,3,4-trifluorophenyl)methyl]thio]- (9CI) (CA INDEX NAME)

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RN 356057-90-4 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(2-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-91-5 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-N-methyl-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-92-6 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[2-(1-piperidinyl)ethyl]-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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eH₂-s-N

RN 356057-93-7 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[3-[4-(trifluoromethyl)phenoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

RN 356057-94-8 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4-[4-(trifluoromethyl)phenoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-95-9 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[3-[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]propyl]- (9CI) (CA INDEX NAME)

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CN 1H-Cyclopentapyrimidine-1-acetamide, N-(1-ethyl-4-piperidinyl)-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356057-99-3 CAPLUS

CN

1H-Cyclopentapyrimidine-1-acetamide, N-[2-(ethylamino)-2-methylpropyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356058-00-9 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-[(1,1-dimethylethyl)amino]ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356058-03-2 CAPLUS

CN Carbamic acid, [2-[[[2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-1H-cyclopentapyrimidin-1-yl]acetyl][[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]amino]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

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RN 356058-05-4 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-N-(1-methyl-4-piperidinyl)-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356058-06-5 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-N-[1-(1-methylethyl)-4-piperidinyl]-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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RN 356058-07-6 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-N-[1-(2-methoxyethyl)-4-piperidinyl]-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{CF3} \\ \\ \text{CH2} \\ \text{N} \\ \text{C} \\ \text{C$$

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RN 356058-12-3 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(ethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

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L3 29 S L1 SSS FUL

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